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Proposal for a Revised Classification of the Depth of Neuromuscular Block and Suggestions for Further Development in Neuromuscular Monitoring

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Anesthesiologists are experiencing new demands for the induction and maintenance of deep neuromuscular block for certain surgical interventions and simultaneously witnessing technical advances in neuromuscular monitoring equipment. The authors view this article as an educational proposal in a process that has gained new momentum and that addresses both clinicians and the manufacturers of medical devices.

The discussion whether a deep or even total neuromuscular block might be justified for surgeries such as robot-assisted endoscopic, ophthalmological, or airway operations is still ongoing. Some authors state that maintaining a deep neuromuscular block between 1 and 3 post-tetanic counts during pneumoperitoneum or operations in the posterior chamber of the eye is mandatory.^{1,2} The authors support their claim with the well-accepted patient safety argument that inadvertent movements of insufficiently paralyzed patients can cause significant morbidity. It is self-explanatory that this cannot be proven in prospective, randomized studies. Fernando et al³ have demonstrated that even at the very low post-tetanic count value of 1, the diaphragm still can inadvertently contract. If this happens, endoscopic instruments can injure the intra-abdominal organs, large vessels, or other delicate structures. Another argument favoring deep block is the improved access to the operation field in the abdominal cavity by the surgeons, if the abdominal wall is completely relaxed.^{4,5} The application of lower insufflation pressures of carbon dioxide into the abdominal cavity during a very deep block may cause less postoperative pain associated with residual carbon dioxide below the diaphragm.^{6,7} Even if these findings remain controversial, we believe there is a need to redefine the nomenclature for the spectrum of neuromuscular block. With a

clearer differentiation of deep block levels, we might facilitate future discussions.

Naguib et al⁸ have proposed the following stratification of the neuromuscular block spectrum (Table 1), which is widely accepted. We consider the designated area for “deep block” in Naguib list, which ranges from a train-of-four count = 0 to a post-tetanic count ≥ 1 to be too broad. Although this classification may be sufficient for the majority of surgical purposes, for those who prefer to work with the very deep and narrow segment of post-tetanic counts values ranging from 1 to 3, this area should be specifically identified. For this particular range, we propose the term “profound block.” Thus, our slightly modified scale divides the original deep block as shown in Table 2.

When extubating a patient’s trachea, the difference between “minimal” and “shallow” block is important. We believe that a distinction between a deep and profound block is also justified, considering their significance for specific surgeries. We must emphasize that the proposal for introducing the profound block level into this list of definitions is not meant to encourage clinicians to achieve deeper block levels than they otherwise would. However, for some surgical procedures, such as intraocular, where even minor patient movements may be disastrous, and when diaphragmatic contractions must be prevented to avoid increases in intracranial pressure associated with tracheal suctioning, profound levels of block are recommended.^{3,9} Ultimately, these are decisions that each clinician must make based on individual clinical need. Our proposal merely fine-tunes the set of definitions of the various depths of neuromuscular block in an attempt to standardize the terminology. In addition, although profound block may be necessary in certain settings to prevent injury, the clinician cannot and must not assume that this technique is devoid of significant side effects. First, many clinicians do not have unrestricted access to sugammadex, while neuromuscular antagonism from profound degrees of block is ineffective with neostigmine. Second, the very surgeries that may require profound block (laparoscopic, robotic, eye, and airway surgery) have surgical closure times that are much shorter than traditional open abdominal procedures. Therefore, the depth of block at the time of neostigmine reversal may be deeper than of other surgeries, rendering patients at increased risk of residual neuromuscular block or recurrence of block (“recurarization”).

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Table 1. Original Neuromuscular Block Levels by Naguib et al⁸

| Level of Block | Depth of Block | Objective Measurement at the Adductor Pollicis | Subjective Measurement at the Adductor Pollicis |
|----------------|---------------------|---|---|
| Level 5 | Complete block | Post-tetanic count = 0 | Post-tetanic count = 0 |
| Level 4 | Deep block | Post-tetanic count ≥ 1 , train-of-four count = 0 | Post-tetanic count ≥ 1 , train-of-four count = 0 |
| Level 3 | Moderate block | Train-of-four count = 1–3 | Train-of-four count = 1–3 |
| Level 2b | Shallow block | Train-of-four ratio < 0.4 | Train-of-four count = 4; train of four fade present |
| Level 2a | Minimal block | Train-of-four ratio = 0.4–0.9 | Train-of-four count = 4; train of four fade is not detectable |
| Level 1 | Acceptable recovery | Train-of-four ratio ≥ 0.9 | Cannot be determined |

Levels of neuromuscular block. Subjective evaluation of neuromuscular block is not recommended, but it is included as an interim transition from current practice to the preferred, objective monitoring-based practice.

Table 2. Modified Neuromuscular Block Levels

| Level of Block | Depth of Block | Objective Measurement at the Adductor Pollicis | Subjective Measurement at the Adductor Pollicis |
|----------------|---------------------|---|---|
| Level 7 | Complete block | Post-tetanic count = 0 | Post-tetanic count = 0 |
| Level 6 | Profound block | Post-tetanic count = 1–3 | Post-tetanic count = 1–3 |
| Level 5 | Deep block | Post-tetanic count ≥ 4 , train-of-four count = 0 | Post-tetanic count ≥ 4 , train-of-four count = 0 |
| Level 4 | Moderate block | Train-of-four count = 1–3 | Train-of-four count = 1–3 |
| Level 3 | Shallow block | Train-of-four ratio < 0.4 | Train-of-four count = 4; train of four fade present |
| Level 2 | Minimal block | Train-of-four ratio = 0.4–0.9 | Train-of-four count = 4; train of four fade is not detectable |
| Level 1 | Acceptable recovery | Train-of-four ratio ≥ 0.9 | Cannot be determined |

Adapted from Naguib et al⁸ with renamed levels of block and split former deep block level and integer level numbering.

A second topic we believe needs to be addressed is the display of neuromuscular monitoring data over time. Older technology, such as the TOF-Watch (Organon, Ireland), allowed the user to download the intraoperative data onto an interfaced computer and display them in a trend format; these data included train-of-four ratio, baseline twitch height, train-of-four count, skin temperature, etc. Modern neuromuscular monitors should have a built-in trend function that can be reviewed by the clinician contemporaneously and should have the ability to be annotated by the user. We would welcome equipment with implemented algorithms that modify both the stimulation pattern and the interval time according to the result of the last measurement. Newer neuromuscular monitoring equipment such as the TOFscan (IdMed, Marseille, France), TetraGraph (Senzime BV, Uppsala, Sweden), and TOFcuff (RGB Medical, Madrid, Spain) display time and dosing-related courses of neuromuscular block with the main stimulation patterns of train-of-four ratio, train-of-four count, and post-tetanic counts. In some of these devices, a semiautomatic or even a fully automatic mode is already implemented.

However, this is still not sufficient. In an ideal system, during anesthesia and neuromuscular block induction, the device would measure TOF in short intervals (eg, every 20 seconds) as long as a train-of-four count is present. As soon as train-of-four count becomes 0 (which might be confirmed by a second measurement), the device should assess the post-tetanic counts. This would indicate that the neuromuscular block level has reached a deep block, where TOF stimulation does not yield positive values. Because of the more pronounced release of synaptic acetylcholine induced by the tetanic stimulation, the time interval after a post-tetanic count sequence should then automatically switch to a longer period (eg, 3 or 4 minutes) and avoid measurements during post-tetanic potentiation.¹⁰ In addition, before each post-tetanic count measurement, a TOF count of zero (train-of-four count = 0) should ensure that a tetanic

stimulation is only delivered during deep or profound block. This sequence of train-of-four count = 0 followed by a post-tetanic count sequence should be repeated every 3 minutes until train-of-four count becomes positive (> 0), in which case the sequence should automatically revert to train-of-four count every 20 seconds until train-of-four ratio > 0.9 . Users could set threshold limits for a desired neuromuscular block level, which would deliver a warning when a measurement is outside of the desired range.

Concerning the monitor screen, we suggest using intuitive symbols on a large screen, which enables an overview of a longer period, preferably with a variable time scale resolution. In the Figure, the consecutive display of the entire spectrum of neuromuscular block values is illustrated, as implemented to various degrees in the TetraGraph and in an experimental version of the TOFcuff software (Figure). In addition to these clear and self-explanatory symbols, we would like to include the option for the user to set a mark that indicates the moment when a repeated dose of neuromuscular block agent has been administered. This could be, for example, a small triangle at the upper screen margin, as depicted in the image. By viewing the neuromuscular blocking pattern and the times of drug dosing, the individual kinetics of a patient can be better understood, thus leading to better individualized neuromuscular blocking agent dosing.

A better delineation of the levels of intraoperative neuromuscular block, an improved trend display of the varying depths of intraoperative block with appropriate symbols to indicate important events, and the appearance on the market of new quantitative monitors are all significant events that should improve patient care.

Summarizing our proposals, we would modify Naguib scale by dividing the original deep block into profound block (post-tetanic counts = 1–3) and deep block (post-tetanic count ≥ 4 and train-of-four count = 0). We also suggest renumbering the levels with integers. New neuromuscular

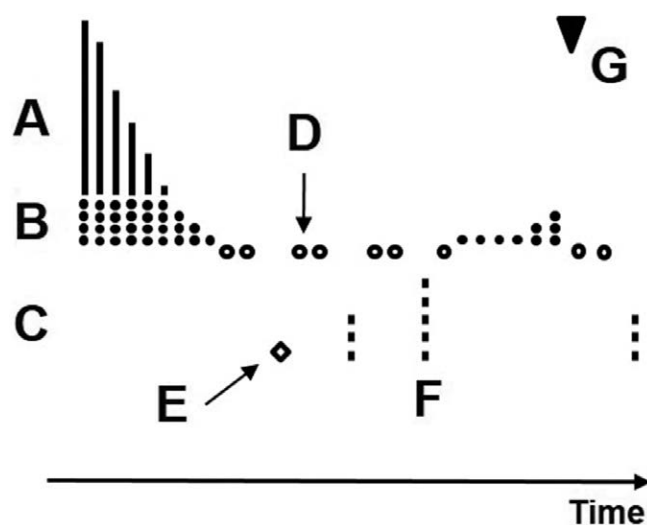


Figure. Symbols indicating the responses to various stimulation patterns displayed over time. The image shows an example of the succession of values that may occur after an initial bolus dose of a nondepolarizing neuromuscular blocker and after 1 repetition dose. A, Fading train-of-four ratio (TOFR) after induction. B, Number of prevailing and receding train-of-four count (TOFC). C, Region for dots representing post-tetanic count (PTC) values. D, Symbol for TOFC = 0. E, Symbol for PTC = 0. F, A PTC value of 5. The triangle on the top right corner (G) is user set and indicates the time when a repetitive dose of neuromuscular blocking agent has been administered. The different time intervals between the measurements are not displayed proportionally in this image.

monitoring equipment may implement an automatic mode that adapts to the stimulation pattern and the time intervals to the measured values. Finally, a larger portion of the neuromuscular block course would be visible if displayed on a screen in a landscape orientation or as a module on the monitoring display, thus illustrating the timely context of dose and effect in a clear and intuitive fashion. ■■

DISCLOSURES

Name: Peter Biro, MD, DESA.

Contribution: This author helped voice the necessity to present this topic and with the original idea for the frame and structure of the manuscript, and helped create the figure and write most of the text.

Conflicts of Interest: P. Biro has given recommendations for the configuration of the display in the TOFcuff monitor and is medical adviser of Guide In Medical, Nazareth, Israel, and has received travel allowances from Merck Sharp & Dohme AG, Switzerland.

Name: Georgina Paul, MD, MBChB.

Contribution: This author helped write parts of the text and completed the language editing and final check.

Conflicts of Interest: None.

Name: Albert Dahan, MD, PhD.

Contribution: This author helped discuss and solve the controversies and write the parts of the text.

Conflicts of Interest: A. Dahan has received consultancy/speaker fees from MSD Nederland BV as well as research funding for projects of his department at LUMC. In addition, he received consultancy fees from Medtronic and Medasense.

Name: Sorin J. Brull, MD, FCARCSI.

Contribution: This author helped discuss and solve the controversies in light of the cited paper by Naguib et al,⁸ and helped write parts of the text, and make a final check.

Conflicts of Interest: S. J. Brull has intellectual property assigned to Mayo Clinic (Rochester, MN); has received research funding from Merck & Co, Inc (funds to Mayo Clinic) and is a consultant for Merck & Co, Inc (Kenilworth, NJ); is a principal and shareholder in Senzime AB (publ) (Uppsala, Sweden); is a contributor to UpToDate (Wolters Kluwer, Waltham, MA); and a member of the Scientific Advisory Boards for ClearLine MD (Woburn, MA), The Doctors Company (Napa, CA), and NMD Pharma (Aarhus, Denmark).

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